

Liquid dosage forms for oral administration can include pharmaceutically acceptable emulsions, solutions, suspensions, syrups, and elixirs containing inert diluents commonly used in the art, such as water.

- 5 Such compositions can also comprise adjuvants, such as wetting agents, emulsifying and suspending agents, and sweetening, flavoring, and perfuming agents.

The amount of active ingredient that can be combined with the carrier materials to produce a single  
10 dosage form varies depending upon the mammalian host treated and the particular mode of administration.

#### Dosage of COX-2 Inhibitors

Dosage levels of COX-2 inhibitors on the order of  
15 about 0.1 mg to about 10,000 mg of the active antiangiogenic ingredient compound are useful in the treatment of the above conditions, with preferred levels of about 1.0 mg to about 1,000 mg. The amount of active  
20 ingredient that may be combined with other anticancer agents to produce a single dosage form will vary depending upon the host treated and the particular mode of administration.

It is understood, however, that a specific dose level for any particular patient will depend upon a  
25 variety of factors including the activity of the specific compound employed, the age, body weight, general health, sex, diet, time of administration, rate of excretion, drug combination, and the severity of the particular disease being treated and form of  
30 administration.

Treatment dosages generally may be titrated to optimize safety and efficacy. Typically, dosage-effect

relationships from in vitro initially can provide useful guidance on the proper doses for patient administration. Studies in animal models also generally may be used for guidance regarding effective dosages for treatment of

5 cancers in accordance with the present invention. In terms of treatment protocols, it should be appreciated that the dosage to be administered will depend on several factors, including the particular agent that is administered, the route administered, the condition of

10 the particular patient, etc. Generally speaking, one will desire to administer an amount of the compound that is effective to achieve a serum level commensurate with the concentrations found to be effective in vitro. Thus, where an compound is found to demonstrate in vitro

15 activity at, e.g., 10  $\mu$ M, one will desire to administer an amount of the drug that is effective to provide about a 10  $\mu$ M concentration in vivo. Determination of these parameters are well within the skill of the art. These considerations, as well as effective formulations and

20 administration procedures are well known in the art and are described in standard textbooks.

The phrase "antineoplastic agents" includes agents that exert antineoplastic effects, i.e., prevent the development, maturation, or spread of neoplastic cells,

25 directly on the tumor cell, e.g., by cytostatic or cytocidal effects, and not indirectly through mechanisms such as biological response modification. There are large numbers of antineoplastic agents available in commercial use, in clinical evaluation and in pre-

30 clinical development, which could be included in the present invention for treatment of neoplasia by combination drug chemotherapy. For convenience of

discussion, antineoplastic agents are classified into the following classes, subtypes and species:

- ACE inhibitors,
- alkylating agents,
- 5 angiogenesis inhibitors,
- angiostatin,
- anthracyclines/DNA intercalators,
- anti-cancer antibiotics or antibiotic-type agents,
- antimetabolites,
- 10 antimetastatic compounds,
- asparaginases,
- bisphosphonates,
- cGMP phosphodiesterase inhibitors,
- calcium carbonate,
- 15 cyclooxygenase-2 inhibitors
- DHA derivatives,
- DNA topoisomerase,
- endostatin,
- epipodophylotoxins,
- 20 genistein,
- hormonal anticancer agents,
- hydrophilic bile acids (URSO),
- immunomodulators or immunological agents,
- integrin antagonists
- 25 interferon antagonists or agents,
- MMP inhibitors,
- miscellaneous antineoplastic agents,
- monoclonal antibodies,
- nitrosoureas,
- 30 NSAIDs,
- ornithine decarboxylase inhibitors,
- pBATTs,